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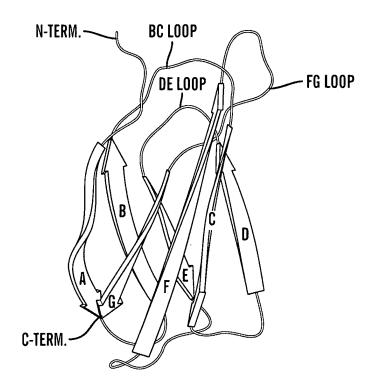


FIG. 1A

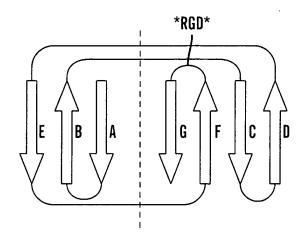
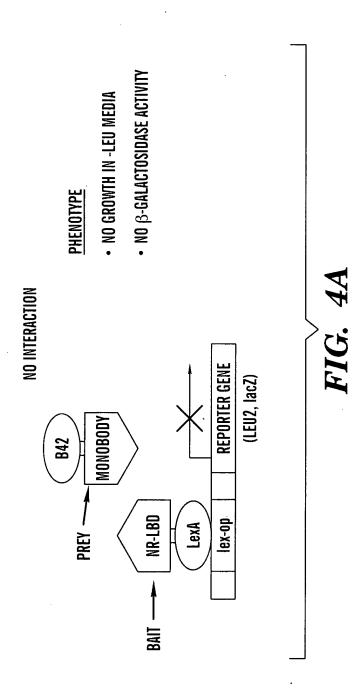
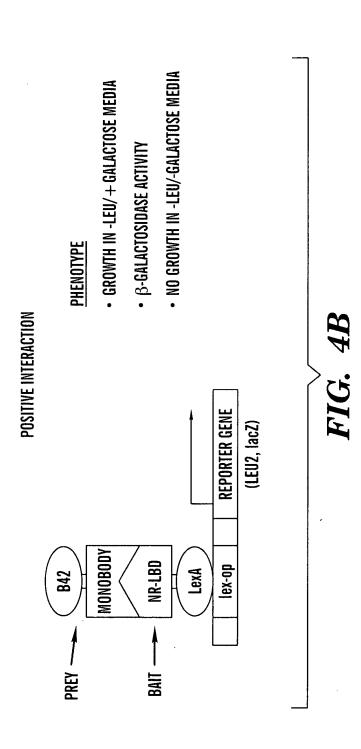


FIG. 1B

NdeI			
CATATGCAGGTTTCTGATGTTCCGCG			
MetGlnValSerAspValProAr	gAspLeuGluVa	lValAlaAlaThrPr	coThrSer
-2 -1 1	. 1	.0	
		_	
BclI PvuII PstI	3.0000.000		BsiWI
CTGCTGATCAGCTGGGATGCTCCTGC			
LeuLeuIleSerTrpAspAlaProAl			rTyrGLy
20	3	0	
	EcoRI		
GAAACCGGTGGTAACTCCCCGGTTCA		' ል ሮሮሞሮሮሞሞሮሮል ልፎሞር	יייזערייכריי
GluThrGlyGlyAsnSerProValGl			
40		illioorybernysse 0	TIHLALO
••	J		
Sa	lI .	Bst1107I	
ACCATCAGCGGCCTGAAACCGGGTGT	CGACTATACCAT	CACTGTATACGCTGT	TACTGGC
ThrIleSerGlyLeuLysProGlyVa			1 1
60	7	0	L
SacI			
XhoI			
CGTGGTGACAGCCCAGCGAGCTCCAA			GTAACTC
ArgGlyAspSerProAlaSerSerLy			
80	9	0	
BamHI			
GAGGATCC			
ONCORTO		,	
•	•		

ECORI		EcoRI	
Psti 31 41 SWDAPAVTVR YYRITYGETG GNSPVQEFTV C D	Xhoi 91 SI NYRT G	Pstl 31 41 SWDAPAVTVR YYRITYGETG GNSPVQEFTV C D	XhoI 91 SI NYRT G
31 YYRITYGE C	Saci 91 SPASSKPISI NYRT	31 YYRITYGE C	Saci 81 91 SPASSKP <u>ISI NYRT</u>
	Salı 71 KPGV \overline{DYTI} TVYAV \overline{TGRGD} \overline{F} $FIG.~3A$. 0	PGSKSTATIS GLKPGVDYTI TVYAVTGRGD FIG. 3B
11 VAATPTSLLI A B	Salı 61 5 GLKPGV <u>DYT</u> 1	11 VAATPTSLLI A B	Sall 61 GLKPGV <u>DYT</u> I
Ndel 1 1 11 MQ VSDVPRDLEV VAATPTSLLI A B	51 PGSK <u>STATI</u> S GL E	Ndel 1 11 MQ VSDVPRXLEV VAATPTSLLI A B	51 PGSK <u>STATI</u> S
MQ		Ndei 1 MQ V	





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PvuII

GAAGTTGTTGCTGCGACCCCGACTAGCCTGCTGATCAGCTGGGATGCTCCT**NNKNNKNNK** GluValValAlaAlaThrProThrSerLeuLeuIleSerTrpAspAlaProXaaXaaXaa

EcoRI

NNKNNKTATTACCGTATCACGTACGGTGAAACCGGTGGTAACTCCCCGGTTCAGGAATTC XaaXaaTyrTyrArgIleThrTyrGlyGluThrGlyGlyAsnSerProValGlnGluPhe

SalI

ACTGTACCTGGTTCCAAGTCTACTGCTACCATCAGCGGCCTGAAACCGGGTGTCGACTAT ThrValProGlySerLysSerThrAlaThrIleSerGlyLeuLysProGlyValAspTyr

ACCATCACTGTATACGCTGTTACTGGC**NNKNNKNNKNNKNNKNNKNNK**TCCAAGCCAATC ThrIleThrValTyrAlaValThrGlyXaaXaaXaaXaaXaaXaaSerLysProIle

KpnI

TCGATTAACTACCGTACCAGTGGTACCGGTGGTTCCCCTCCAAAAAAGAAGAAGAAAGGTA SerIleAsnTyrArgThrSerGlyThrGlyGlySerProProLysLysArgLysVal

GCTGGTATCAATAAAGATATCGAGGAGTGCAATGCCATCATTGAGCAGTTTATCGACTAC AlaGlyIleAsnLysAspIleGluGluCysAsnAlaIleIleGluGlnPheIleAspTyr

CTGCGCACCGGACAGGAGATGCCGATGGAAATGGCGGATCAGGCGATTAACGTGGTGCCGLeuArgThrGlyGlnGluMetProMetGluMetAlaAspGlnAlaIleAsnValValPro

GGCATGACGCCGAAAACCATTCTTCACGCCGGGCCGCCGATCCAGCCTGACTGGCTGAAA GlyMetThrProLysThrIleLeuHisAlaGlyProProIleGlnProAspTrpLeuLys

TCGAATGGTTTTCATGAAATTGAAGCGGATGTTAACGATACCAGCCTCTTGCTGAGTGGA SerAsnGlyPheHisGluIleGluAlaAspValAsnAspThrSerLeuLeuLeuSerGly

XhoI SphI GATTAACTCGAGGCATGC Asp•••

FIG. 5

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ATGGGTAAGCCTATCCCTAACCCTCTCCTCGGTCTCGATTCTACACAAGCTATGGGTGCT MetGlyLysProIleProAsnProLeuLeuGlyLeuAspSerThrGlnAlaMetGlyAla

CCTCCAAAAAAGAGAGAAAGGTAGCTGGTATCAATAAAGATATCGAGGAGTGCAATGCC ProProLysLysLysArgLysValAlaGlyIleAsnLysAspIleGluGluCysAsnAla

ATCATTGAGCAGTTTATCGACTACCTGCGCACCGGACAGGAGATGCCGATGGAAATGGCGIleIleGluGlnPheIleAspTyrLeuArgThrGlyGlnGluMetProMetGluMetAla

GATCAGGCGATTAACGTGGTGCCGGGCATGACGCCGAAAACCATTCTTCACGCCGGGCCG AspGlnAlaIleAsnValValProGlyMetThrProLysThrIleLeuHisAlaGlyPro

CCGATCCAGCCTGACTGGCTGAAATCGAATGGTTTTCATGAAATTGAAGCGGATGTTAAC ProlleGlnProAspTrpLeuLysSerAsnGlyPheHisGluIleGluAlaAspValAsn KpnI

HindIII SacI

GATACCAGCCTCTTGCTGAGTGGAGATGCCTCCAAGCTTGGTACCGAGCTCGGATCTATG AspThrSerLeuLeuSerGlyAspAlaSerLysLeuGlyThrGluLeuGlySerMet

CAGGTTTCTGATGTTCCGACCGACCTGGAAGTTGTTGCTGCGACCCCG**NNSNNSNNSNNS** GlnValSerAspValProThrAspLeuGluValValAlaAlaThrProXaaXaaXaaXaa

PvuII PstI

NNSNNSACTAGCCTGCTGATCAGCTGGGATGCTCCTGCAGTTACCGTGCGTTATTAC XaaXaaXaaThrSerLeuLeuIleSerTrpAspAlaProAlaValThrValArgTyrTyr

EcoRI

CGTATCACGTACGGTGAAACCGGTGGTAACTCCCCGGTTCAGGAATTCACTGTACCTGGT ArgIleThrTyrGlyGluThrGlyGlyAsnSerProValGlnGluPheThrValProGly

SalI

TCCAAGTCTACTGCTACCATCAGCGGCCTGAAACCGGGTGTCGACTATACCATCACTGTA SerLysSerThrAlaThrIleSerGlyLeuLysProGlyValAspTyrThrIleThrVal

SacI

TACGCTGTTACTGGCCGTGGTGACAGCCCAGCGAGCTCCAAGCCAATCTCGATTAACTAC
TyrAlaValThrGlyArgGlyAspSerProAlaSerSerLysProIleSerIleAsnTyr

XhoI SphI CGTACCTAGTAACTCGAGGCATGC ArgThr•••••

FIG. 6

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ATGGGTAAGCCTATCCCTAACCCTCTCCTCGGTCTCGATTCTACACAAGCTATGGGTGCT MetGlyLysProIleProAsnProLeuLeuGlyLeuAspSerThrGlnAlaMetGlyAla

CCTCCAAAAAAGAGAGAAAGGTAGCTGGTATCAATAAAGATATCGAGGAGTGCAATGCC ProProLysLysLysArgLysValAlaGlyIleAsnLysAspIleGluGluCysAsnAla

 $\label{lem:attacca} \textbf{ATCATTGAGCAGTTATCGACTACCTGCGCACCGGACAGGAGATGCCGATGGAAATGGCGIleIleGluGlnPheIleAspTyrLeuArgThrGlyGlnGluMetProMetGluMetAla}$

GATCAGGCGATTAACGTGGTGCCGGGCATGACGCCGAAAACCATTCTTCACGCCGGGCCG AspGlnAlaIleAsnValValProGlyMetThrProLysThrIleLeuHisAlaGlyPro

CCGATCCAGCCTGACTGGCTGAAATCGAATGGTTTTCATGAAATTGAAGCGGATGTTAAC ProIleGlnProAspTrpLeuLysSerAsnGlyPheHisGluIleGluAlaAspValAsn KpnI

HindIII SacI

GATACCAGCCTCTTGCTGAGTGGAGATGCCTCCAAGCTTGGTACCGAGCTCGGATCTATG AspThrSerLeuLeuLeuSerGlyAspAlaSerLysLeuGlyThrGluLeuGlySerMet

CAGGTTTCTGATGTTCCGACCGACCTGGAAGTTGTTGCTGCGACCCCGACTAGCCTGCTGGInValSerAspValProThrAspLeuGluValValAlaAlaThrProThrSerLeuLeu

PvuII

ATCAGCTGGGATGCTCCT**NNKNNKNNKNNK**TATTACCGTATCACGTACGGTGAAACC IleSerTrpAspAlaProXaaXaaXaaXaaXaaTyrTyrArgIleThrTyrGlyGluThr

EcoRI

SalI

AGCGGCCTGAAACCGGGTGTCGACTATACCATCACTGTATACGCTGTTACTGGC**NNKNNK**SerGlyLeuLysProGlyValAspTyrThrIleThrValTyrAlaValThrGlyXaaXaa

XhoI SphI

NNKNNKNNKNNKTCCAAGCCAATCTCGATTAACTACCGTACCTAGTAACTCGAGGCA XaaXaaXaaXaaXaaSerLysProIleSerIleAsnTyrArgThr••••••

TGCATCTAGAGGGCCGCATCATGTAATTAGTTATGTCACGCTTA

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 $\label{thm:condition} \textbf{ATGGGTAAGCCTATCCTCGGTCTCGATTCTACACAAGCTATGGGTGCT} \\ \textbf{MetGlyLysProIleProAsnProLeuLeuGlyLeuAspSerThrGlnAlaMetGlyAla} \\$

CCTCCAAAAAAGAGAGAAAGGTAGCTGGTATCAATAAAGATATCGAGGAGTGCAATGCC ProProLysLysArgLysValAlaGlyIleAsnLysAspIleGluGluCysAsnAla

 $\label{lem:attacca} ATCATTGAGCAGTTATCGACTACCTGCGCACCGGACAGGAGATGCCGATGGAAATGGCGIleIleGluGlnPheIleAspTyrLeuArgThrGlyGlnGluMetProMetGluMetAla$

GATCAGGCGATTAACGTGGTGCCGGGCATGACGCCGAAAACCATTCTTCACGCCGGGCCG AspGlnAlaIleAsnValValProGlyMetThrProLysThrIleLeuHisAlaGlyPro

CCGATCCAGCCTGACTGGCTGAAATCGAATGGTTTTCATGAAATTGAAGCGGATGTTAAC
ProlleGlnProAspTrpLeuLysSerAsnGlyPheHisGluIleGluAlaAspValAsn
KpnI

HindIII SacI

 ${\tt GATACCAGCCTCTTGCTGAGTGGAGATGCCTCCAAGCTTGGTACCGAGCTCGGATCTATGASpThrSerLeuLeuSerGlyAspAlaSerLysLeuGlyThrGluLeuGlySerMet}$

CGTGTTTCTGATGTTCCGCGTGACCTGGAAGTTGTTGCTGCGACCCCGACTAGCCTGCTG ArgValSerAspValProArgAspLeuGluValValAlaAlaThrProThrSerLeuLeu

PvuTT

ATCAGCTGGGATGCTCCTGCAGTTACCGTGCGTTATTACCGTATCACGTACGGTGAAACC IleSerTrpAspAlaProAlaValThrValArgTyrTyrArgIleThrTyrGlyGluThr

EcoRI

 $\label{thm:condition} GGTGGTAACTCCCGGTTCAGGAATTCACTGTACCTGGTTCCAAGTCTACTGCTACCATC \\ GlyGlyAsnSerProValGlnGluPheThrValProGlySerLysSerThrAlaThrIle\\$

SalI

AGCGGCCTGAAACCGGGTGTCGACTATACCATCACTGTATACGCTGTTACTGGC**NNKNNK** SerGlyLeuLysProGlyValAspTyrThrIleThrValTyrAlaValThrGlyXaaXaa

XhoI SphI TACCGTACCTAGTAACTCGAGGCATGC TyrArgThr••••••

FIG. 8

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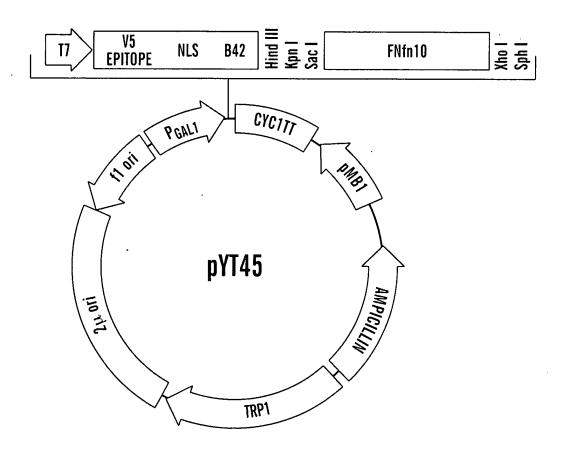


FIG. 9

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ATGGGTAAGCCTATCCCTAACCCTCTCCTCGGTCTCGATTCTACACAAGCTATGGGTGCT MetGlyLysProIleProAsnProLeuLeuGlyLeuAspSerThrGlnAlaMetGlyAla

CCTCCAAAAAAGAGAGAAAGGTAGCTGGTATCAATAAAGATATCGAGGAGTGCAATGCC ProProLysLysArgLysValAlaGlyIleAsnLysAspIleGluGluCysAsnAla

ATCATTGAGCAGTTTATCGACTACCTGCGCACCGGACAGGAGATGCCGATGGAAATGGCG IleIleGluGlnPheIleAspTyrLeuArgThrGlyGlnGluMetProMetGluMetAla

GATCAGGCGATTAACGTGGTGCCGGGCATGACGCCGAAAACCATTCTTCACGCCGGGCCG AspGlnAlaIleAsnValValProGlyMetThrProLysThrIleLeuHisAlaGlyPro

CCGATCCAGCCTGACTGGCTGAAATCGAATGGTTTTCATGAAATTGAAGCGGATGTTAAC ProlleGlnProAspTrpLeuLysSerAsnGlyPheHisGluIleGluAlaAspValAsn

HindIII/KpnI/SacI

GATACCAGCCTCTTGCTGAGTGGAGATGCCTCCAAGCTTGGTACCGAGCTCGGATCTATG AspThrSerLeuLeuLeuSerGlyAspAlaSerLysLeuGlyThrGluLeuGlySerMet

 ${\tt CAGGTTTCTGATGTTCCGACCGACCTGGAAGTTGTTGCTGCGACCCCGACTAGCCTGCTGGInValSerAspValProThrAspLeuGluValValAlaAlaThrProThrSerLeuLeu}$

PvuII PstI

ATCAGCTGGGATGCTCCTGCAGTTACCGTGCGTTATTACCGTATCACGTACGGTGAAACC IleSerTrpAspAlaProAlaValThrValArgTyrTyrArgIleThrTyrGlyGluThr

EcoRI

GGTGGTAACTCCCCGGTTCAGGAATTCACTGTACCTGGTTCCAAGTCTACTGCTACCATC GlyGlyAsnSerProValGlnGluPheThrValProGlySerLysSerThrAlaThrIle

SalI

AGCGGCCTGAAACCGGGTGTCGACTATACCATCACTGTATACGCTGTTACTGGCCGTGGT SerGlyLeuLysProGlyValAspTyrThrIleThrValTyrAlaValThrGlyArgGly

SacI XhoI SphI GACAGCCCAGCGAGCTCCAAGCCAATCTCGATTAACTACCGTACCTAGTAACTCGAGGCA AspSerProAlaSerSerLysProIleSerIleAsnTyrArgThr • • • • • •

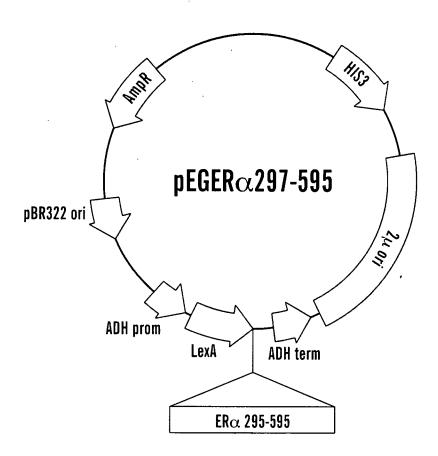


FIG. 11

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ATGAAAGCGTTAACGGCCAGGCAACAAGAGGTGTTTGATCTCATCCGTGATCACATCAGC MetLysAlaLeuThrAlaArgGlnGlnGluValPheAspLeuIleArgAspHisIleSer CAGACAGGTATGCCGCCGACGCGTGCGGAAATCGCGCAGCGTTTGGGGGTTCCGTTCCCCA GlnThrGlyMetProProThrArgAlaGluIleAlaGlnArgLeuGlyPheArgSerPro AACGCGGCTGAAGAACATCTGAAGGCGCTGGCACGCAAAGGCGTTATTGAAATTGTTTCC AsnAlaAlaGluGluHisLeuLysAlaLeuAlaArgLysGlyValIleGluIleValSer GGCGCATCACGCGGGATTCGTCTGTTGCAGGAAGAGGAAGAAGGGTTGCCGCTGGTAGGT GlyAlaSerArgGlyIleArgLeuLeuGlnGluGluGluGluGlyLeuProLeuValGly cgtgtggctgccggtgaaccacttctggcgcaacagcatattgaaggtcattatcaggtc ArgValAlaAlaGlyGluProLeuLeuAlaGlnGlnHisIleGluGlyHisTyrGlnVal GATCCTTCCTTATTCAAGCCGAATGCTGATTTCCTGCTGCGCGTCAGCGGGATGTCGATG AspProSerLeuPheLysProAsnAlaAspPheLeuLeuArgValSerGlyMetSerMet AAAGATATCGGCATTATGGATGGTGACTTGCTGGCAGTGCATAAAACTCAGGATGTACGT LysAspIleGlyIleMetAspGlyAspLeuLeuAlaValHisLysThrGlnAspValArg AACGGTCAGGTCGTTGTCGCACGTATTGATGACGAAGTTACCGTTAAGCGCCTGAAAAAA AsnGlyGlnValValValAlaArgIleAspAspGluValThrValLysArgLeuLysLys CAGGGCAATAAAGTCGAACTGTTGCCAGAAAATAGCGAGTTTAAACCAATTGTCGTAGAT GlnGlyAsnLysValGluLeuLeuProGluAsnSerGluPheLysProIleValValAsp CTTCGTCAGCAGAGCTTCACCATTGAAGGGCTGGCGGTTGGGGGTTATTCGCAACGGCGAC LeuArgGlnGlnSerPheThrIleGluGlyLeuAlaValGlyValIleArgAsnGlyAsp SacI

EcoRI HindIII

AGCCTGGCCTTGTCCCTGACGGCCGACCAGATGGTCAGTGCCTTGTTGGATGCTGAGCCC SerLeuAlaLeuSerLeuThrAlaAspGlnMetValSerAlaLeuLeuAspAlaGluPro

HindIII

CCCATACTCTATTCCGAGTATGATCCTACCAGACCCTTCAGTGAAGCTTCGATGATGGGC ProlleLeuTyrSerGluTyrAspProThrArgProPheSerGluAlaSerMetMetGly

FIG. 12A

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TTACTGACCAACCTGGCAGACAGGGAGCTGGTTCACATGATCAACTGGGCGAAGAGGGTG LeuLeuThrAsnLeuAlaAspArgGluLeuValHisMetIleAsnTrpAlaLysArgVal

XbaI

CCAGGCTTTGTGGATTTGACCCTCCATGATCAGGTCCACCTTCTAGAATGTGCCTGGCTA ProGlyPheValAspLeuThrLeuHisAspGlnValHisLeuLeuGluCysAlaTrpLeu

GAGATCCTGATGATTGGTCTCGTCTGGCGCTCCATGGAGCACCCAGTGAAGCTACTGTTTGluIleLeuMetIleGlyLeuValTrpArgSerMetGluHisProValLysLeuLeuPhe

GCTCCTAACTTGCTCTTGGACAGGAACCAGGGAAAATGTGTAGAGGGCATGGTGGAGATC AlaProAsnLeuLeuAspArgAsnGlnGlyLysCysValGluGlyMetValGluIle

PstI

TTTGTGTGCCTCAAATCTATTATTTTGCTTAATTCTGGAGTGTACACATTTCTGTCCAGC PheValCysLeuLysSerIleIleLeuLeuAsnSerGlyValTyrThrPheLeuSerSer

ACCCTGAAGTCTCTGGAAGAGAAGACCATATCCACCGAGTCCTGGACAAGATCACAGAC ThrLeuLysSerLeuGluGluLysAspHisIleHisArgValLeuAspLysIleThrAsp

PstI

ACTTTGATCCACCTGATGGCCAAGGCAGCAGCAGCAGCAGCAGCAGCAGCAGCTG ThrLeuIleHisLeuMetAlaLysAlaGlyLeuThrLeuGlnGlnHisGlnArgLeu

GCCCAGCTCCTCCTCCTCCCCACATCAGGCACATGAGTAACAAAGGCATGGAGCAT AlaGlnLeuLeuIleLeuSerHisIleArgHisMetSerAsnLysGlyMetGluHis

CTGTACAGCATGAAGTGCAAGAACGTGGTGCCCCTCTATGACCTGCTGCTGGAGATGCTG LeuTyrSerMetLysCysLysAsnValValProLeuTyrAspLeuLeuGluMetLeu

GACGCCCACCGCCTACATGCGCCCACTAGCCGTGGAGGGGCATCCGTGGAGGAGACGGACASpAlaHisArgLeuHisAlaProThrSerArgGlyGlyAlaSerValGluGluThrAsp

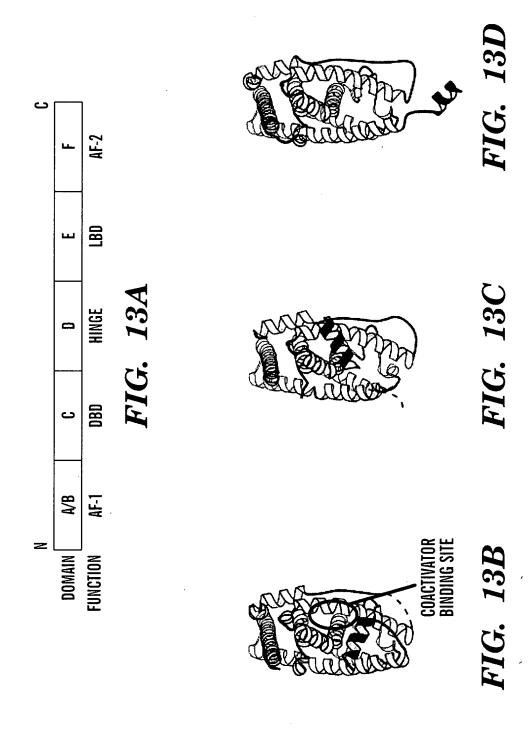
CAAAGCCACTTGGCCACTGCGGGCTCTACTTCATCGCATTCCTTGCAAAAGTATTACATC GlnSerHisLeuAlaThrAlaGlySerThrSerSerHisSerLeuGlnLysTyrTyrIle

XhoI

ACGGGGGAGGCAGAGGGTTTCCCTGCCACAGTCTGACtcgag ThrGlyGluAlaGluGlyPheProAlaThrVal•••

FIG. 12B

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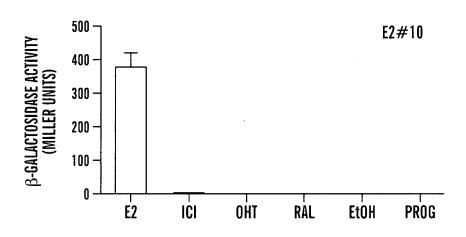


FIG. 14A

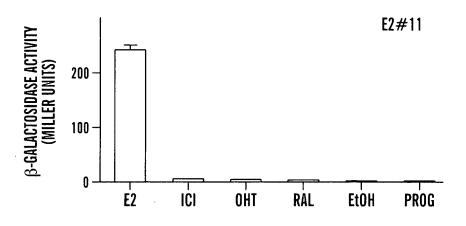


FIG. 14B

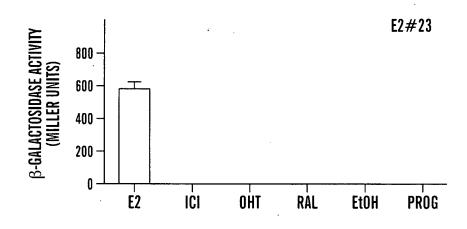


FIG. 14C

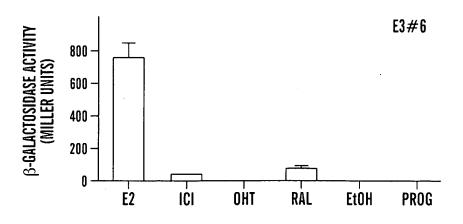


FIG. 14D

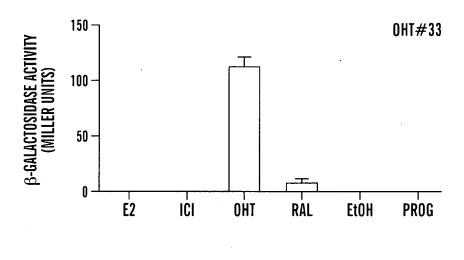


FIG. 14E

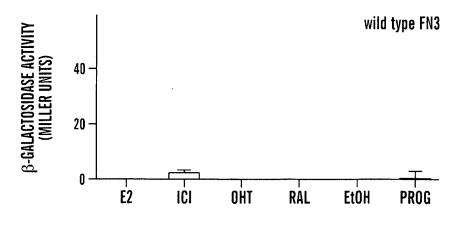


FIG. 14F

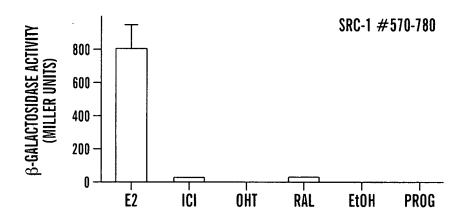


FIG. 14G

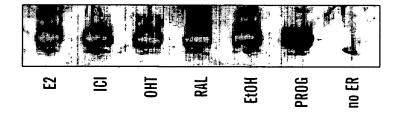


FIG. 14H

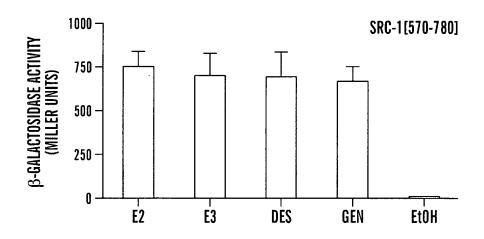


FIG. 15A

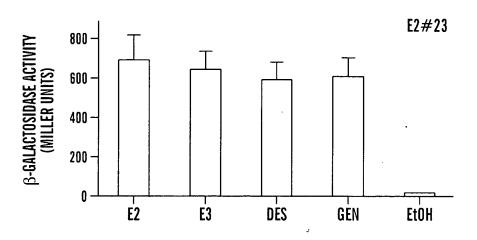


FIG. 15B

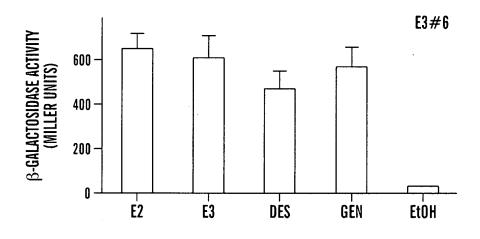


FIG. 15C

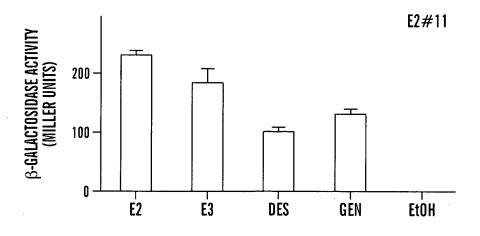


FIG. 15D

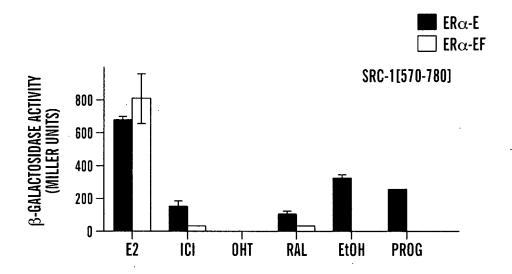


FIG. 16A

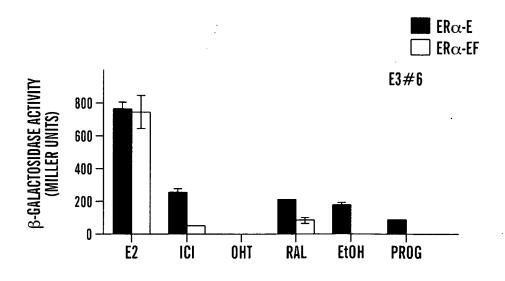
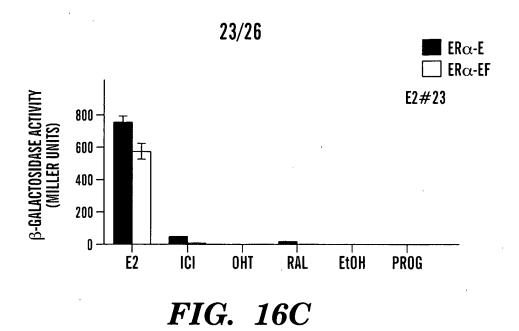
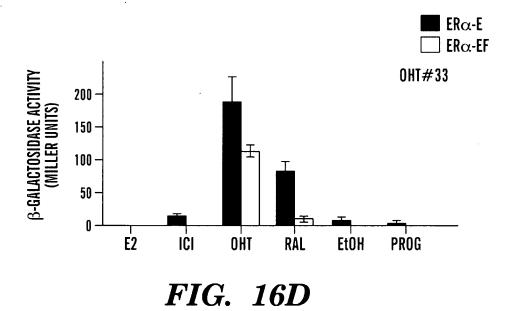
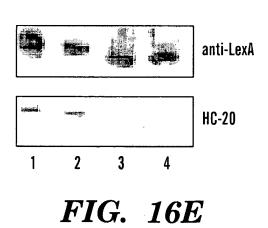


FIG. 16B







NEW SHEET

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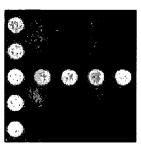
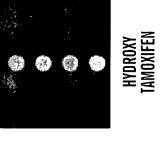


FIG. 17D



ESTRADIOL

FIG. 17C

FIG. 17B

FIG. 17A

NO SELECTION (MASTER PLATE)

NO LIGAND

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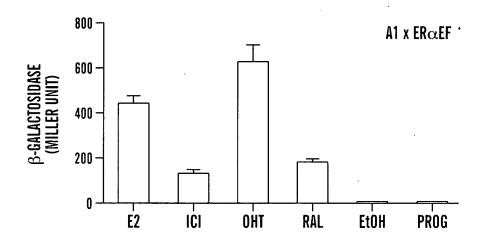


FIG. 18A

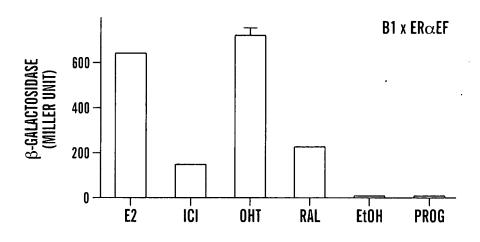


FIG. 18B

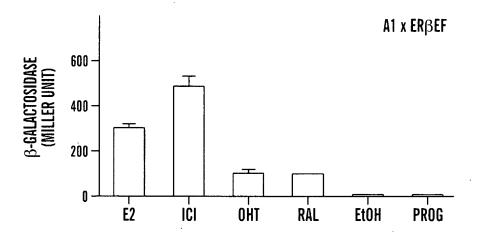


FIG. 18C

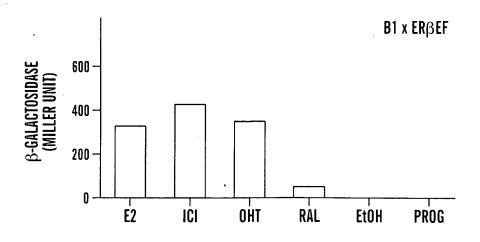


FIG. 18D